(1) GENERAL INFORMATION

(ii) TITLE OF INVENTION: ANTI-PHOSPHORYLATED TAU PROTEIN ANTIBODY AND METHOD FOR DETECTING ALZHEIMER'S DISEASE WITH THE USE OF THE SAME

(iii) NUMBER OF SEQUENCES: 22

(iv) COMPUTER READABLE FORM:

- (A) MEDIUM TYPE: Floppy disk
- (B) COMPUTER: IBM PC compatible
- (C) OPERATING SYSTEM: PC-DOS/MS-DOS

Kazutomo Imahori

- (D) SOFTWARE: PatentIn Release #1.0, Version #1.30 (EPO)
- (vi) CURRENT APPLICATION DATA
 - (A) APPLICATION NUMBER:
 - (B)FILING DATE:
 - (C)CLASSIFICATION:
- (vi) PRIOR APPLICATION DATA
 - (A)APPLICATION NUMBER: JP 8/56090
 - (B)FILING DATE: 13-MAR-1996
- (2) INFORMATION FOR SEQ ID NO: 1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 441 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY:/linear
 - (ii) MOLECULE TYPE ≠ protein
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

Met Ala Glu Pro Arg Gin Glu Phe Glu Val Met Glu Asp His Ala Gly

1 5 / 10 15

Gln Asp Thr Tyr Gly/Leu Gly Asp Arg Lys Asp Gln Gly Gly Tyr Thr
20 25 30

Met His Gln Glu Gly Asp Thr Asp Ala Gly Leu Lys Glu Ser Pro Leu

3	5		40					45			
Gln Thr Pr	o Thr Glu	Asp Gly	ser Ser	Glu	Glu	Pro	Gly	Ser	Glu	Thr	Ser
50		55					60				
Asp Ala Ly	s Ser Thi		Ala	Glu	Asp		Thr	Ala	Pro	Leu	Val
65		70			_	75					80
Asp Glu Gl	y Ala Pro. 85		GIn	Ala	90	Ala	Gln	Pro	His	Thr 95	Glu
Ile Pro Gl	u Gly Thi 100	Thr Ala	Glu	Glu 105	Ala	Gly	Ile	Gly	Asp 110	Thr	Pro
Ser Leu Gl	_	ı Ala Ala	Gly 120	His	Val	Thr	Gln	Ala 125	Arg	Met	Val
Ser Lys Se	r Lys Asp	Gly Thr		Ser	Asp	Asp	Lys 140	Lys	Ala	Lys	Gly
Ala Asp Gl	y Lys Thi			Thr	Pro	Arg	Gly	Ala	Ala	Pro	Pro
145		150				155	_				160
Gly Gln Ly	s Gly Glr 165		Ala	Thr	Arg 170	Ile	Pro	Ala	Lys	Thr 175	Pro
Pro Ala Pr	o Lys Thr	Pro Pro	Ser	Ser	Gly	Glu	Pro	Pro	Lys	Ser	Gly
	180			185					190		
Asp Arg Se		Ser Ser		Gly	Ser	Pro	Gly		Pro	Gly	Ser
19			200	1	_	_	_,	205		_	
Arg Ser Ar 210	g Thr Pro	ser Lev 215		Thr	Pro	Pro	1hr 220	Arg	GLu	Pro	Lys
Lys Val Al	a Val Val			Pro	Twe	Ser		Ser	Ser	Δla	Tage
225	a var var	230	110	110	11,5	235	110	DCL	DCI	ALG	240
Ser Arg Le	u Gln Thr		Val	Pro	Met		Asp	Leu	Lys	Asn	
	245	i			250		_		_	255	
Lys Ser Ly	s Ile Gly	Ser Thr	Glu	Asn	Leu	Lys	His	Gln	Pro	Gly	Gly
	260			265					270		
Gly Lys Va		: Ile Asn		Lys	Leu	Asp	Leu		Asn	Val	Gln
27		. T	280	T1.	T	TT	17-7	285 December 2	01	a 1.	a i
Ser Lys Cy 290	s Gly Sei	Lys Asp 295		тте	гàг	HIS	300	Pro	GTĀ	GIĄ	GTÀ
Ser Val Gl	n Ile Val		Pro	Val	Asp	Leu	Ser	Lys	Val	Thr	
305		310			•	315					320
Lys Cys Gl	y Ser Leu 325		ı Ile		His 330	Lys	Pro	Gly	Gly	Gly 335	Gln
Val Glu Va		Glu Lys		_	Phe	Lys	Asp	Arg		Gln	Ser
T T	340	•		345	 ·		_		350		
Lys Ile Gl	y ser Leu	ı Asp Asn	ıııe	'I'nr	Hls	va⊥	Pro	GLY	GLy	GLy	Asn

355 360 Lys Lys Ile Glu Thr His Lys Leu Thr Phe Arg Glu Asn Ala Lys Ala 370 375 380 Lys Thr Asp His Gly Ala Glu Ile Val Tyr Lys Ser Pro Val Val Ser 390 395 400 Gly Asp Thr Ser Pro Arg His Leu Ser Asn Val Ser Ser Thr Gly Ser 405 410 415 Ile Asp Met Val Asp Ser Pro Gln Leu Ala Thr Leu Ala Asp Glu Val 430 Ser Ala Ser Leu Ala Lys Gln Gly Leu 435 440

- (2) INFORMATION FOR SEQ ID NO: 2:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 6
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:
 Lys Ser Gly Tyr Ser Xaa Pro Gly Ser Pro Gly Thr
 1 5 10
- (2) INFORMATION FOR SEQ ID NO: 3:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 13 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 6
 - (D) OTHER INFORMATION: Xaa = phosphoserine

- (2) INFORMATION FOR SEQ ID NO: 4:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphothreonine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4: Cys Pro Gly Ser Pro Gly Xaa Pro Gly Ser Arg Ser

1 5 10

- (2) INFORMATION FOR SEQ ID NO: 5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 13 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 6
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

 Lys Ser Xaa Pro Gly Xaa Pro Gly Thr Pro Gly Ser Arg

 1 5 10
- (2) INFORMATION FOR SEQ ID NO: 6:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 14 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphothreonine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:
 Cys Val Ala Val Val Arg Xaa Pro Pro Lys Ser Pro Ser Ser

- (2) INFORMATION FOR SEQ ID NO: 7:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7: Cys Arg Thr Pro Pro Lys Xaa Pro Ser Ser Ala Lys 1 5 10
- (2) INFORMATION FOR SEQ ID NO: 8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (2) INFORMATION FOR SEQ ID NO: 9:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 3
 - (D) OTHER INFORMATION: Xaa = phosphothreonine

And of other in the state of th

- (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:

Cys Arg Xaa Pro Pro Lys Xaa Pro Ser Ser Ala Lys 5

1

10

- (2) INFORMATION FOR SEQ ID NO: 10:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:

Cys Lys Ser Lys Ile Gly Xaa Thr Glu Asn Leu Lys

1

(2) INFORMATION FOR SEQ ID NO: 11:

5

- (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

Cys Glu Ile Val Tyr Lys Xaa Pro Val Val Ser Gly

1

5

10

10

- (2) INFORMATION FOR SEQ ID NO: 12:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

- (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12: Cys Val Ser Gly Asp Thr Xaa Pro Arg His Leu Ser
- (2) INFORMATION FOR SEQ ID NO: 13:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:
 Lys Leu Ser Asn Val Ser Xaa Thr Gly Ser Ile Asp
 1 5 10
- (2) INFORMATION FOR SEQ ID NO: 14:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (2) INFORMATION FOR SEQ ID NO: 15:
 - (i) SEQUENCE CHARACTERISTICS:

- (A) SEQUENCE LENGTH: 12 amino acids(B) SEQUENCE TYPE: amino acid
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 6
 - (D) OTHER INFORMATION:Xaa=phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

 Lys Leu Ser Asn Val Xaa Ser Thr Gly Ser Ile Asp

 1 5 10
- (2) INFORMATION FOR SEQ ID NO: 16:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 6
 - (D) OTHER INFORMATION: Xaa = phosphoserine
 - (ix) FEATURE:
 - (A) NAME/KEY:
 - (B) LOCATION: 7
 - (D) OTHER INFORMATION: Xaa = phosphoserine
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16: Lys Leu Ser Asn Val Xaa Xaa Thr Gly Ser Ile Asp

1 5 10

- (2) INFORMATION FOR SEQ ID NO: 17:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 17 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

Ser Pro Gln Leu Ala Thr Leu Ala Asp Glu Val Ser Ala Ser Leu Ala Lys

1 5

10

	-49-									
(2)	<pre>INFORMATION FOR SEQ ID NO: 18: (i) SEQUENCE CHARACTERISTICS: (A) SEQUENCE LENGTH: 12 amino acids (B) SEQUENCE TYPE: amino acid (D) TOPOLOGY: linear</pre>									
	(ii) MOLECULE TYPE: peptide									
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:									
Ala	Glu Pro Arg Gln Glu Phe Glu Val Met Glu Cys									
1	5 10									
(2)	<pre>INFORMATION FOR SEQ ID NO: 19: (i) SEQUENCE CHARACTERISTICS: (A) SEQUENCE LENGTH: 15 amino acids (B) SEQUENCE TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:</pre>									
Val	Ala Val Val Arg Thr Pro Pro Lys Ser Pro Ser Ser Ala Lys									
1	5 10 15									
(2)	<pre>INFORMATION FOR SEQ ID NO: 20: (i) SEQUENCE CHARACTERISTICS: (A) SEQUENCE LENGTH: 34 amino acids (B) SEQUENCE TYPE: amino acid (D) TOPOLOGY: linear</pre>									
	(ii) MOLECULE TYPE: peptide									
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:									
Ser	Gly Asp Arg Ser Gly Tyr Ser Ser Pro Gly Ser Pro Gly Thr P	ro								
1	5 10 15									
Gly	Ser Arg Ser Arg Thr Pro Ser Leu Pro Thr Pro Pro Thr Arg G	lu								
-	20 25 30									
Pro	rys									
(2)	INFORMATION FOR SEQ ID NO: 21:									

- (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 55 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

Ala Lys Thr Asp His Gly Ala Glu Ile Val Tyr Lys Ser Pro Val Val

1 5 10 15

Ser Gly Asp Thr Ser Pro Arg His Leu Ser Asn Val Ser Ser Thr Gly
20 25 30

Ser Ile Asp Met Val Asp Ser Pro Gln Leu Ala Thr Leu Ala Asp Glu 35 40 45

Val Ser Ala Ser Leu Ala Lys 50 55

- (2) INFORMATION FOR SEQ ID NO: 22:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) SEQUENCE LENGTH: 12 amino acids
 - (B) SEQUENCE TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide

• / •

the first three first in the first the first three the first three first three